

In the Claims

The listing of claims will replace all prior versions and listings of claims in the application.

Listings of claims

Claim 1. (original) A combination, comprising an endothelin receptor antagonist, or a pharmaceutically acceptable salt thereof, and an EGFR TKI, or a pharmaceutically acceptable salt thereof.

Claim 2. (original) A combination according to claim 1 wherein the endothelin receptor antagonist is selected from A-127722, atrasentan (ABT-627), BQ-123, BQ-788, BMS 182874, feloprentan, BSF 420627, FR139317, IPI-950, L-749,329, L-754,142, LU 110896, LU 110897, PD 156707, PD 155080, Ro 46-2005, bosentan (Ro 47-0203), SB 217242, SB 209670, TAK-044, YM598, sitaxsentan (TBC11251), ambrisentan, tezosentan, darusentan, *N*-[[2'-[[4,5-dimethyl-3-isoxazolyl]amino]sulphonyl]-4-(2-oxazolyl)[1,1'-biphenyl]-2-yl]methyl]-N,3,3-trimethylbutanamide, ZD1611 and *N*-(3-methoxy-5-methylpyrazin-2-yl)-2-(4-[1,3,4-oxadiazol-2-yl]phenyl)pyridine-3-sulphonamide (ZD4054), or a pharmaceutically acceptable salt thereof.

Claim 3. (currently amended) A combination according to claim 1 [[or 2]] wherein the EGFR TKI is selected from:

N-(3-chloro-4-fluorophenyl)-7-methoxy-6-(3-morpholinopropoxy)quinazolin-4-amine (ZD1839);

N-(3-ethynylphenyl)-6,7-bis(2-methoxyethoxy)quinazolin-4-amine, or a pharmaceutically-acceptable salt thereof (linked to the code numbers CP 358774 and OSI-774 (the monomethanesulphonate salt));

6-acrylamido-*N*-(3-chloro-4-fluorophenyl)-7-(3-morpholinopropoxy)quinazolin-4-amine (linked to the code numbers PD 183805 and CI 1033);

4-[(1R)-1-phenylethylamino]-6-(4-hydroxyphenyl)-7H-pyrrolo[2,3-*d*]pyrimidine (linked to the code numbers PKI-166, CGP 75166 and CGP 59326);

N-[4-(3-bromoanilino)quinazolin-6-yl]but-2-ynamide (linked to the code numbers CL-387785 and EKB-785); and

4-(3-chloro-4-fluoroanilino)-3-cyano-6-(4-dimethylaminobut-2(E)-enamido)-7-ethoxyquinoline (EKB-569);

or a pharmaceutically acceptable salt thereof.

Claim 4. (currently amended) A combination according to ~~any one of claims 1-3~~ claim 1 wherein the endothelin receptor antagonist is selected from ZD4054, or a pharmaceutically acceptable salt thereof, and the EGFR TKI is selected from ZD1839, or a pharmaceutically acceptable salt thereof.

Claim 5. (cancelled)

Claim 6. (currently amended) A pharmaceutical composition comprising a combination according to ~~any one of claim[[s]] 1[-4]~~, in association with a pharmaceutically acceptable diluent or carrier.

Claim 7. (currently amended) A method of treating cancer, in a warm-blooded animal, such as man, in need of such treatment which comprises administering to said animal an effective amount of a combination according to ~~any one of claim[[s]] 1[-4]~~.

Claim 8. - 9. (cancelled)

Claim 10. (currently amended) The method ~~or use or combination~~ according to claim[[s]] 7[[
-9]] wherein the cancer is oesophageal cancer, myeloma, hepatocellular, pancreatic, cervical cancer, ewings tumour, neuroblastoma, kaposi's sarcoma, ovarian cancer, breast cancer, colorectal cancer, prostate cancer, bladder cancer, melanoma, lung cancer - non small cell lung cancer (NSCLC), and small cell lung cancer (SCLC), gastric cancer, head and neck cancer, brain cancer, renal cancer, lymphoma and leukaemia.

Claim 11. (cancelled)

Claim 12. (currently amended) The method ~~or use or combination~~ according to claim 10 wherein the cancer is prostate cancer.

Claim 13. (currently amended) The method ~~or use or combination~~ according to claim 10 wherein the cancer is NSCLC.

Claim 14. (currently amended) The method ~~or use or combination~~ according to claim 10 wherein the cancer is in a metastatic state.

Claim 15. (currently amended) The method or use or combination according to claim 10 wherein the cancer is in a non-metastatic state.

Claim 16. (currently amended) The method or use or combination according to claim 10 wherein the cancer is renal, thyroid, lung, breast or prostate cancer that is producing bone metastases.

Claim 17. (new) A combination according to claim 2 wherein the EGFR TKI is selected from:

N-(3-chloro-4-fluorophenyl)-7-methoxy-6-(3-morpholinopropoxy)quinazolin-4-amine (ZD1839);

N-(3-ethynylphenyl)-6,7-bis(2-methoxyethoxy)quinazolin-4-amine, or a pharmaceutically-acceptable salt thereof (linked to the code numbers CP 358774 and OSI-774 (the monomethanesulphonate salt));

6-acrylamido-*N*-(3-chloro-4-fluorophenyl)-7-(3-morpholinopropoxy)quinazolin-4-amine (linked to the code numbers PD 183805 and CI 1033);

4-[(1*R*)-1-phenylethylamino]-6-(4-hydroxyphenyl)-7*H*-pyrrolo[2,3-*d*]pyrimidine (linked to the code numbers PKI-166, CGP 75166 and CGP 59326);

N-[4-(3-bromoanilino)quinazolin-6-yl]but-2-ynamide (linked to the code numbers CL-387785 and EKB-785); and

4-(3-chloro-4-fluoroanilino)-3-cyano-6-(4-dimethylaminobut-2(*E*)-enamido)-7-ethoxyquinoline (EKB-569);

or a pharmaceutically acceptable salt thereof.

Claim 18. (new) A pharmaceutical composition comprising a combination according to claim 17, in association with a pharmaceutically acceptable diluent or carrier.

Claim 19. (new) A method of treating cancer, in a warm-blooded animal, such as man, in need of such treatment which comprises administering to said animal an effective amount of a combination according to claim 17.

Claim 20. (new) The method according to claim 19 wherein the cancer is prostate cancer.

Claim 21. (new) The method according to claim 19 wherein the cancer is NSCLC.

Claim 22. (new) The method according to claim 19 wherein the cancer is in a metastatic state.

Claim 23. (new) The method according to claim 19 wherein the cancer is in a non-metastatic state.

Claim 24. (new) The method according to claim 19 wherein the cancer is renal, thyroid, lung, breast or prostate cancer that is producing bone metastases.